```
                10        20        30        40        50        60
HUMAN   MGIVEPGCGDMLTGTEPMPGSDEGRAPGADPQHRYFYPEPGAQDADERRGGGSLGSPYPG
        :::::::::::::::::: :::::.:::: :::.::::::::::  .::.:.:::.:: :
MOUSE   MGIVEPGCGDMLTGTEPMP-SDEGRGPGADQQHRFFYPEPGAQDPTDRRAGSSLGTPYSG
                10        20        30        40        50

CONS    MGIVEPGCGDMLTGTEPMP SDEGR PGAD QHR FYPEPGAQD   RR G SLG PY G


                70        80        90       100       110       120
HUMAN   GALVPAPPSRFLGAYAYPPRPQAAGFPGAGESFPPPADAEGYQPGEGYAAPDPRAGLYPG
        :::::: :.::::..::::: :.:::::: :: ::::: :::: : .:: ::::::::::::
MOUSE   GALVPAAPGRFLGSFAYPPRAQVAGFPGPGEFFPPPAGAEGYPPVDGYPAPDPRAGLYPG
      60        70        80        90       100       110

CONS    GALVPA P RFLG  AYPPR Q AGFPG GE FPPPA AEGY P  GY APDPRAGLYPG


               130       140       150       160       170       180
HUMAN   PREDYALPAGLEVSGKLRVALNNHLLWSKFNQHQTEMIITKQGRRMFPFLSFTVAGLEPT
        :::::::::::::::::::::.::::::::::::::::::::::::::::::::::::::
MOUSE   PREDYALPAGLEVSGKLRVALSNHLLWSKFNQHQTEMIITKQGRRMFPFLSFTVAGLEPT
     120       130       140       150       160       170

CONS    PREDYALPAGLEVSGKLRVAL NHLLWSKFNQHQTEMIITKQGRRMFPFLSFTVAGLEPT


               190       200       210       220       230       240
HUMAN   SHYRMFVDVVLVDQHHWRYQSGKWVQCGKAEGSMPGNRLYVHPDSPNTGAHWMRQEVSFG
        ::::::::::::::::::::::::::::::::::::::::::::::::::::::::::::::
MOUSE   SHYRMFVDVVLVDQHHWRYQSGKWVQCGKAEGSMPGNRLYVHPDSPNTGAHWMRQEVSFG
     180       190       200       210       220       230

CONS    SHYRMFVDVVLVDQHHWRYQSGKWVQCGKAEGSMPGNRLYVHPDSPNTGAHWMRQEVSFG


               250       260       270       280       290       300
HUMAN   KLKLTNNKGASNNVTQMIVLQSLHKYQPRLHIVEVNDGEPEAACNASNTHIFTFQETQFI
        ::::::::::::::::::::::::::::::::::::::::::::::.:::::.:::::::::
MOUSE   KLKLTNNKGASNNVTQMIVLQSLHKYQPRLHIVEVNDGEPEAACSASNTHVFTFQETQFI
     240       250       260       270       280       290

CONS    KLKLTNNKGASNNVTQMIVLQSLHKYQPRLHIVEVNDGEPEAAC ASNTH FTFQETQFI
```

Fig. 1A

```
                310        320        330        340        350        360
HUMAN   AVTAYQNAEITQLKIDNNPFAKGFRENFESMYTSVDTSIPSPPGPNCQFLGGDHYSPLLP
        :::::::::::::::::::::::::::::::::::::.:::::.::::::::::.:::: .:::::
MOUSE   AVTAYQNAEITQLKIDNNPFAKGFRENFESMYASVDTSVPSPPGPNCQLLGGDPFSPLLS
      300        310        320        330        340        350

CONS    AVTAYQNAEITQLKIDNNPFAKGFRENFESMY SVDTS PSPPGPNCQ LGGD FSPLL


                370        380        390        400        410        420
HUMAN   NQYPVPSRFYPDLPGQAKDVVPQAYWLGAPRDHSYEAEFRAVSMKPAFLPSAPGPTMSYY
        ::::::::::::::::: ::.. : ::::.::.:::::::::::::..:::::::::. ::
MOUSE   NQYPVPSRFYPDLPGQPKDMISQPYWLGTPREHSYEAEFRAVSMKPTLLPSAPGPTVPYY
      360        370        380        390        400        410

CONS    NQYPVPSRFYPDLPGQ KD   Q YWLG PR HSYEAEFRAVSMKP  LPSAPGPT  YY


                430        440        450        460        470        480
HUMAN   RGQEVLAPGAGWPVAPQYPPKMGPASWFRPMRTLPMEPGPGGSEGRGPEDQGPPLVWTEI
        :::.:::::::::::::::::::.::.:::::::::::.:: :.:: .:    . : .: :.
MOUSE   RGQDVLAPGAGWPVAPQYPPKMSPAGWFRPMRTLPMDPGLGSSEEQG----SSPSLWPEV
      420        430        440        450        460            470

CONS    RGQ VLAPGAGWPVAPQYPPKM PA WFRPMRTLPM PG G SE  G      P  W E


                490        500        510        520        530
HUMAN   APIRPESSDSGLGEGDSKRRRVSPYPSSGDSSSPAGAPSPFDKEAEGQFYNTFPN
        . ..:: ::::::::::.::::.:::::::::::::::::::::::::.:::::::::
MOUSE   TSLQPEPSDSGLGEGDTKRRRISPYPSSGDSSSPAGAPSPFDKETEGQFYNYFPN
          480        490   ↑↑↑↑ 500        510        520        530

CONS        PE SDSGLGEGD KRRR SPYPSSGDSSSPAGAPSPFDKE EGQFYNYFPN
```

Fig. 1A (continued)

```
                10        20        30        40        50        60
HUMAN ATGGGCATCGTGGAGCCGGGTTGCGGAGACATGCTGACGGGCACCGAGCCGATGCCGGGG
      :::::::::::::::::::: ::::::::::::::::: ::::::::::::::::::   :
MOUSE ATGGGCATCGTGGAGCCGGGCTGCGGAGACATGCTGACCGGCACCGAGCCGATGCC---G
                10        20        30        40        50

                70        80        90       100       110       120
HUMAN  AGCGACGAGGGCCGGGCGCCTGGCGCCGACCCGCAGCACCGCTACTTCTACCCGGAGCCG
       :: :::::::::::::: ::: :: :: :::: ::::: :: : :::::: :: ::::::
MOUSE  AGTGACGAGGGCCGGGGGCCCGGAGCGGACCAACAGCATCGTTTCTTCTATCCCGAGCCG
        60        70        80        90       100       110

               130       140       150       160       170       180
HUMAN  GGCGCGCAGGACGCGGACGAGCGTCGCGGGGGCGGCAGCCTGGGGTCTCCCTACCCGGGG
       ::::: :::::: ::  ::: :: :::: ::  :::::::::::: : :::::: : :::
MOUSE  GGCGCACAGGACCCGACCGATCGCCGCGCAGGTAGCAGCCTGGGGACGCCCTACTCTGGG
       120       130       140       150       160       170

               190       200       210       220       230       240
HUMAN  GGCGCCTTGGTGCCCGCCCCGCCGAGCCGCTTCCTTGGAGCCTACGCCTACCCGCCGCGA
       :::::: ::::::: ::: ::::: : :::::::::::: ::: :::::::::::: ::
MOUSE  GCGCCCTGGTGCCTGCCGCGCCGGGTCGCTTCCTTGGATCCTTCGCCTACCCGCCCCGG
       180       190       200       210       220       230

               250       260       270       280       290       300
HUMAN  CCCCAGGCGGCCGGCTTCCCCGGCGCGGGCGAGTCCTTCCCGCCGCCCGCGGACGCCGAG
        : :::: ::: ::::: :::::  : ::::::: ::::::::::::::::::  :: :::
MOUSE  GCTCAGGTGGCTGGCTTTCCCGGGCCTGGCGAGTTCTTCCCGCCGCCCGCGGGTGCGGAG
       240       250       260       270       280       290

               310       320       330       340       350       360
HUMAN  GGCTACCAGCCGGGCGAGGGCTACGCCGCCCCGGACCCGCGCGCCGGGCTCTACCCGGGG
       ::::::: ::: :  :: :::::: : ::::: :::::::::: :::::::::: :::
MOUSE  GGCTACCCGCCCGTGGATGGCTACCCTGCCCCTGACCCGCGCGCGGGGCTCTACCCAGGG
       300       310       320       330       340       350

               370       380       390       400       410       420
HUMAN  CCGCGTGAGGACTACGCGCTACCCGCGGGACTGGAGGTGTCGGGGAAACTGAGGGTCGCG
       ::::: ::::::::::  : ::::::::  ::::::::: ::::: ::::: ::::::
MOUSE  CCGCGCGAGGACTACGCATTGCCCGCGGGGTTGGAGGTGTCTGGGAAGCTGAGAGTCGCG
       360       370       380       390       400       410
```

Fig. 1B

```
          430        440        450        460        470        480
HUMAN     CTCAACAACCACCTGTTGTGGTCCAAGTTTAATCAGCACCAGACAGAGATGATCATCACC
          :::: :::::::::::::::::::::::: :: ::::::::::::::::::::::::::::
MOUSE     CTCAGCAACCACCTGTTGTGGTCCAAGTTCAACCAGCACCAGACAGAGATGATCATCACT
          420        430        440        450        460        470

          490        500        510        520        530        540
HUMAN     AAGCAGGGACGGCGGATGTTCCCATTCCTGTCATTTACTGTGGCCGGGCTGGAGCCCACC
          ::::: :::::::: :::::::::::::::::: :: :: ::::::::::::::::::::
MOUSE     AAGCAAGGACGGCGAATGTTCCCATTCCTGTCCTTCACCGTGGCCGGGCTGGAGCCCACA
          480        490        500        510        520        530

          550        560        570        580        590        600
HUMAN     AGCCACTACAGGATGTTTGTGGACGTGGTCTTGGTGGACCAGCACCACTGGCGGTACCAG
          ::::: ::::::::::::::::: ::::::::::::::::::::::::::::::::::::
MOUSE     AGCCATTACAGGATGTTTGTGGATGTGGTCTTGGTGGACCAGCACCACTGGCGGTACCAG
          540        550        560        570        580        590

          610        620        630        640        650        660
HUMAN     AGCGGCAAGTGGGTGCAGTGTGGAAAGGCCGAGGGCAGCATGCCAGGAAACCGCCTGTAC
          :::::::::::::::::::::::::::::: :: :::::::::::::: :::::: : ::
MOUSE     AGCGGCAAGTGGGTGCAGTGTGGAAAGGCAGAAGGCAGCATGCCAGGGAACCGCTTATAT
          600        610        620        630        640        650

          670        680        690        700        710        720
HUMAN     GTCCACCCGGACTCCCCCAACACAGGAGCGCACTGGATGCGCCAGGAAGTTTCATTTGGG
          :::::::: :::::::::::::: ::::: ::::::::::::::::::::::::::::::
MOUSE     GTCCACCCAGACTCCCCCAACACCGGAGCCCACTGGATGCGCCAGGAAGTTTCATTTGGG
          660        670        680        690        700        710

          730        740        750        760        770        780
HUMAN     AAACTAAAGCTCACAAACAACAAGGGGGCGTCCAACAATGTGACCCAGATGATTGTGCTC
          :: :::::::::::: :::::::::::::: :::::::::::::::::::::::: :: ::
MOUSE     AAGCTAAAGCTCACCAACAACAAGGGGGCTTCCAACAATGTGACCCAGATGATCGTCCTG
          720        730        740        750        760        770

          790        800        810        820        830        840
HUMAN     CAGTCCCTCCATAAGTACCAGCCCCGGCTGCATATCGTTGAGGTGAACGACGGAGAGCCA
          ::::: ::::: :::::::::::::::::::: ::::: :::::::: :: :::::::::
MOUSE     CAGTCTCTCCACAAGTACCAGCCCCGGCTGCACATCGTGGAGGTGAATGATGGAGAGCCA
          780        790        800        810        820        830
```

Fig. 1B (continued)

```
                850       860       870       880       890       900
HUMAN   GAGGCAGCCTGCAACGCTTCCAACACGCATATCTTTACTTTCCAAGAAACCCAGTTCATT
        ::::: ::::::: ::::: ::::: ::  ::::::::::::::::: ::::::::::::
MOUSE   GAGGCTGCCTGCAGTGCTTCTAACACACACGTCTTTACTTTCCAAGAGACCCAGTTCATT
        840       850       860       870       880       890

                910       920       930       940       950       960
HUMAN   GCCGTGACTGCCTACCAGAATGCCGAGATTACTCAGCTGAAAATTGATAATAACCCCTTT
        :: ::::::::::::::::: :: ::::: :::::::::::::: :: :: ::::::::::
MOUSE   GCAGTGACTGCCTACCAGAACGCAGAGATCACTCAGCTGAAAATCGACAACAACCCCTTT
        900       910       920       930       940       950

                970       980       990      1000      1010      1020
HUMAN   GCCAAAGGATTCCGGGAGAACTTTGAGTCCATGTACACATCTGTTGACACCAGCATCCCC
        :::::::::::::::::::::::::::::::::::: :::::::::: :: ::  :::::
MOUSE   GCCAAAGGATTCCGGGAGAACTTTGAGTCCATGTACGCATCTGTTGATACGAGTGTCCCC
        960       970       980       990      1000      1010

               1030      1040      1050      1060      1070      1080
HUMAN   TCCCCGCCTGGACCCAACTGTCAATTCCTTGGGGGAGATCACTACTCTCCTCTCCTACCC
        :: :: :::::::::::::::::: : :::::::::: : :: ::: ::::: ::: ::
MOUSE   TCGCCACCTGGACCCAACTGTCAACTGCTTGGGGGAGACCCCTTCTCACCTCTTCTATCC
        1020      1030      1040      1050      1060      1070

               1090      1100      1110      1120      1130      1140
HUMAN   AACCAGTATCCTGTTCCCAGCCGCTTCTACCCCGACCTTCCTGGCCAGGCGAAGGATGTG
        ::::::::::::::::::::::: ::::::::::::::::: :::::: : :::::: ::
MOUSE   AACCAGTATCCTGTTCCCAGCCGTTTCTACCCCGACCTTCCAGGCCAGCCCAAGGATATG
        1080      1090      1100      1110      1120      1130

              1150      1160      1170      1180      1190      1200
HUMAN   GTTCCCCAGGCTTACTGGCTGGGGGCCCCCCGGGACCACAGCTATGAGGCTGAGTTTCGA
         :  : ::: :::::::::::::: : :: ::::: ::::: ::::: :: ::::: :::
MOUSE   ATCTCACAGCCTTACTGGCTGGGGACACCTCGGGAACACAGTTATGAAGCGGAGTTCCGA
       1140      1150      1160      1170      1180      1190

              1210      1220      1230      1240      1250      1260
HUMAN   GCAGTCAGCATGAAGCCTGCATTCTTGCCCTCTGCCCCTGGGCCCACCATGTCCTACTAC
        :: :: ::::::::::::  :: :: : :::::::::: ::::::::  :: ::::::::
MOUSE   GCTGTGAGCATGAAGCCCACACTCCTACCCTCTGCCCCGGGGCCCACTGTGCCCTACTAC
       1200      1210      1220      1230      1240      1250
```

Fig. 1B (continued)

```
            1270       1280       1290       1300       1310       1320
HUMAN  CGAGGCCAGGAGGTCCTGGCACCTGGAGCTGGCTGGCCTGTGGCACCCCAGTACCCTCCC
       :: ::::: :: :::::::: :::::::::: ::::: ::::: :: :: ::::: :::
MOUSE  CGGGGCCAAGACGTCCTGGCGCCTGGAGCTGGTTGGCCCGTGGCCCCTCAATACCCGCCC
      1260       1270       1280       1290       1300       1310

            1330       1340       1350       1360       1370       1380
HUMAN  AAGATGGGCCCGGCCAGCTGGTTCCGCCCTATGCGGACTCTGCCCATGGAACCCGGCCCT
       :::::: :::: ::  :::::::::: :: ::::: :::::::::::::: :: ::::
MOUSE  AAGATGAGCCCAGCTGGCTGGTTCCGGCCCATGCGAACTCTGCCCATGGACCCGGGCCTG
      1320       1330       1340       1350       1360       1370

            1390       1400       1410       1420       1430       1440
HUMAN  GGAGGCTCAGAGGGACGGGGACCAGAGGACCAGGGTCCCCCCTTGGTGTGGACTGAGATT
       :::  :::::::: :: :::  :                 :::::: : ::::: ::::: :
MOUSE  GGATCCTCAGAGGAACAGGGCTCCT------------CCCCCTCGCTGTGGCCTGAGGTC
      1380       1390       1400                  1410       1420

            1450       1460       1470       1480       1490       1500
HUMAN  GCCCCCATCCGGCCGGAATCCAGTGATTCAGGACTGGGCGAAGGAGACTCTAAGAGGAGG
        :: :: ::: ::::::  :::: :: :::::::: :::::::::::: ::::::::::
MOUSE  ACCTCCCTCCAGCCGGAGCCCAGCGACTCAGGACTAGGCGAAGGAGACACTAAGAGGAGG
        1430       1440       1450       1460       1470       1480

            1510       1520       1530       1540       1550       1560
HUMAN  CGCGTGTCCCCCTATCCTTCCAGTGGTGACAGCTCCTCCCCTGCTGGGGCCCCTTCTCCT
        :  : :::::::::::::::::::: :::::::::: :: ::::::::::::::::::
MOUSE  AGGATATCCCCCTATCCTTCCAGTGGCGACAGCTCCTCTCCCGCTGGGGCCCCTTCTCCT
        1490       1500       1510       1520       1530       1540

            1570       1580       1590       1600
HUMAN  TTTGATAAGGAAGCTGAAGGACAGTTTTATAACTATTTTCCCAACTGA
       :::::::::::: : ::::: :::::::::: ::::::::::::::
MOUSE  TTTGATAAGGAAACCGAAGGCCAGTTTTATAATTATTTTCCCAACTGA
        1550       1560       1570       1580       1590
```

Fig. 1B (continued)

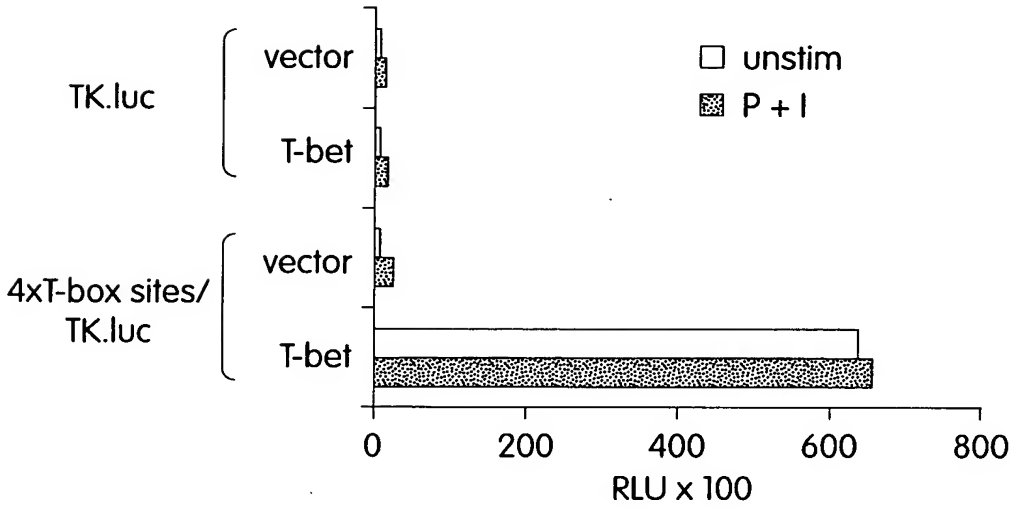
TK.luc
vector
T-bet
4xT-box sites/
TK.luc
vector
T-bet
unstim
P + I
0
200
400
600
800
RLU x 100

Fig. 2A

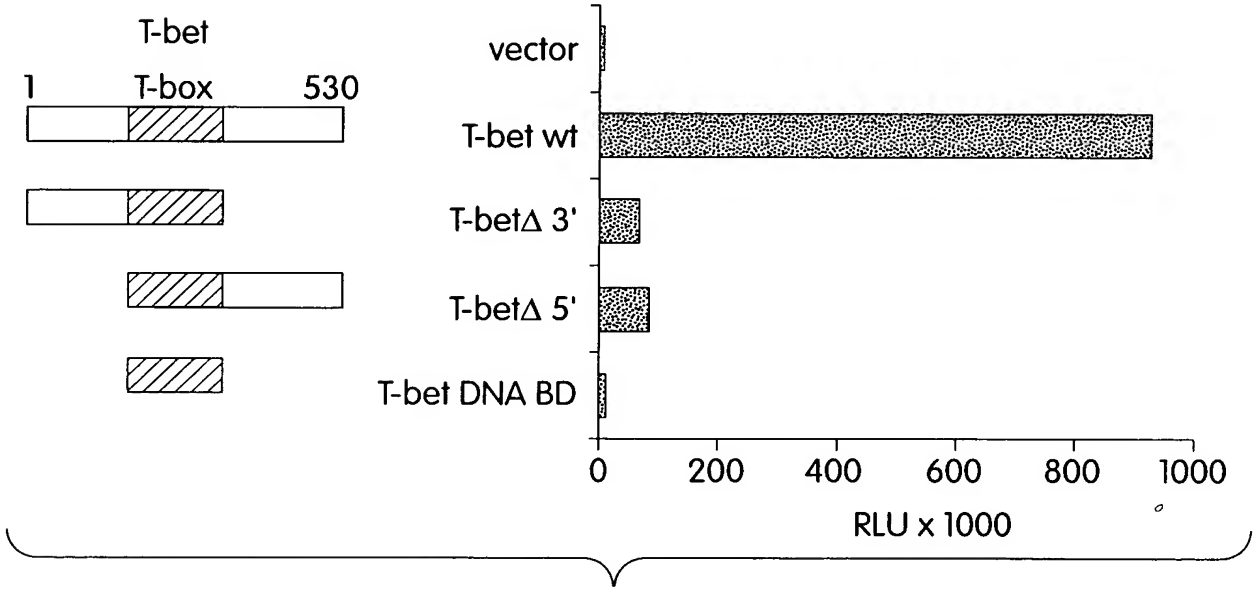
T-bet
1
T-box
530
vector
T-bet wt
T-betΔ 3'
T-betΔ 5'
T-bet DNA BD
0
200
400
600
800
1000
RLU x 1000

Fig. 2B

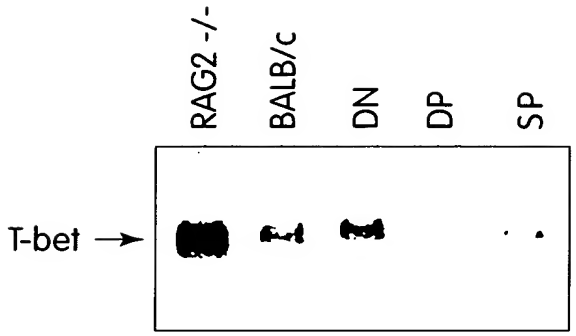
RAG2 -/-
BALB/c
DN
DP
SP
T-bet

Fig. 3A

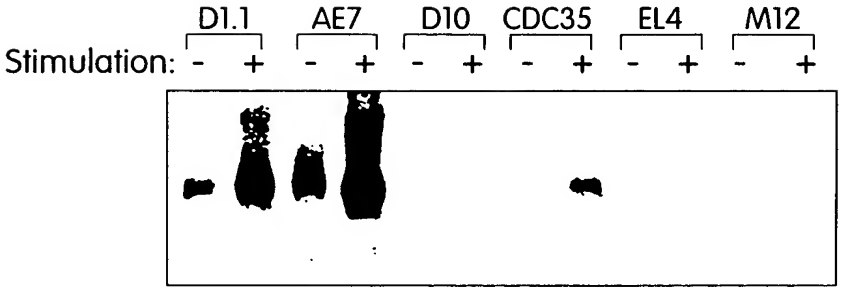
D1.1
AE7
D10
CDC35
EL4
M12
Stimulation: - + - + - + - + - + - +

Fig. 3B

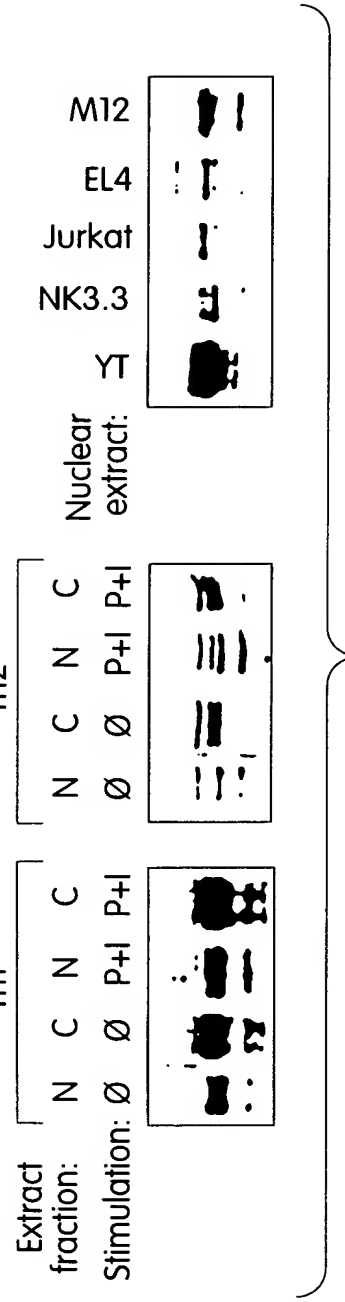
Th1
Th2
Extract fraction:
Stimulation:
N C N C
Ø Ø P+I P+I
N C N C
Ø Ø P+I P+I
Nuclear extract:
M12
EL4
Jurkat
NK3.3
YT

Fig. 3C

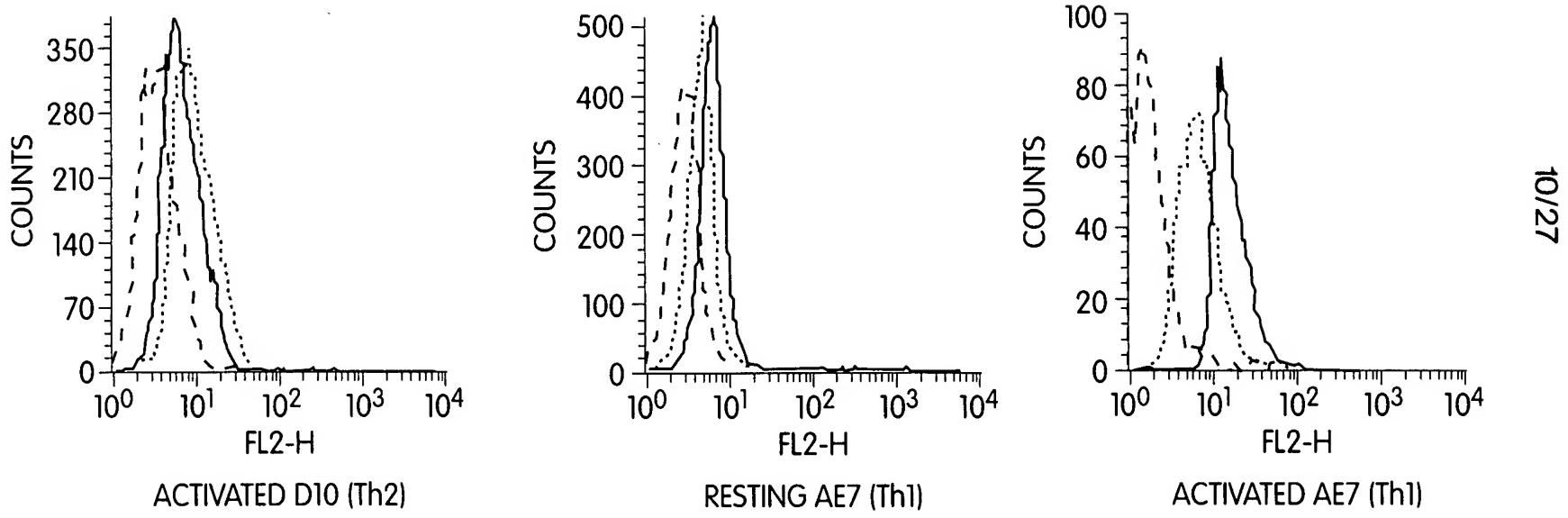
COUNTS
350
280
210
140
70
0
10^0 10^1 10^2 10^3 10^4
FL2-H
ACTIVATED D10 (Th2)
COUNTS
500
400
300
200
100
0
10^0 10^1 10^2 10^3 10^4
FL2-H
RESTING AE7 (Th1)
COUNTS
100
80
60
40
20
0
10^0 10^1 10^2 10^3 10^4
FL2-H
ACTIVATED AE7 (Th1)

Fig. 3D

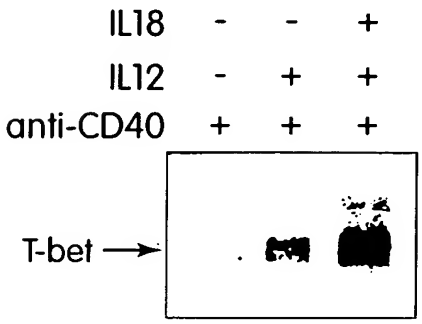

Fig. 4A

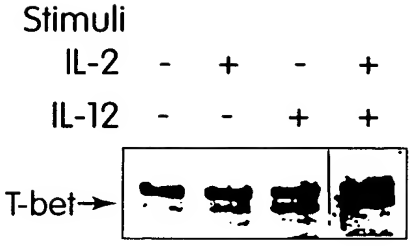

Fig. 4B

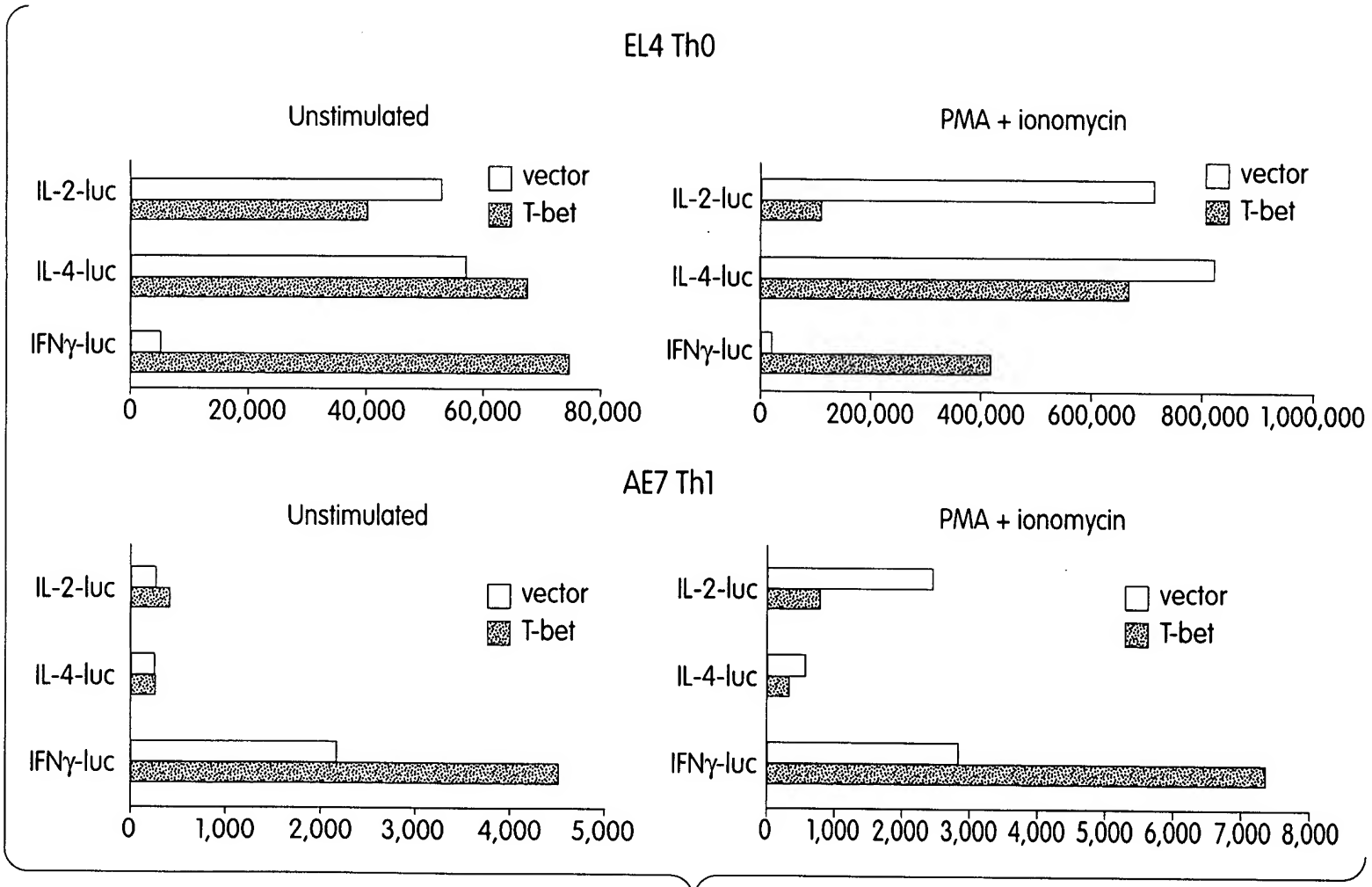
EL4 Th0
Unstimulated
PMA + ionomycin
IL-2-luc
IL-4-luc
IFNγ-luc
vector
T-bet
0
20,000
40,000
60,000
80,000
200,000
400,000
600,000
800,000
1,000,000
AE7 Th1
Unstimulated
PMA + ionomycin
1,000
2,000
3,000
4,000
5,000
6,000
7,000
8,000

Fig. 5

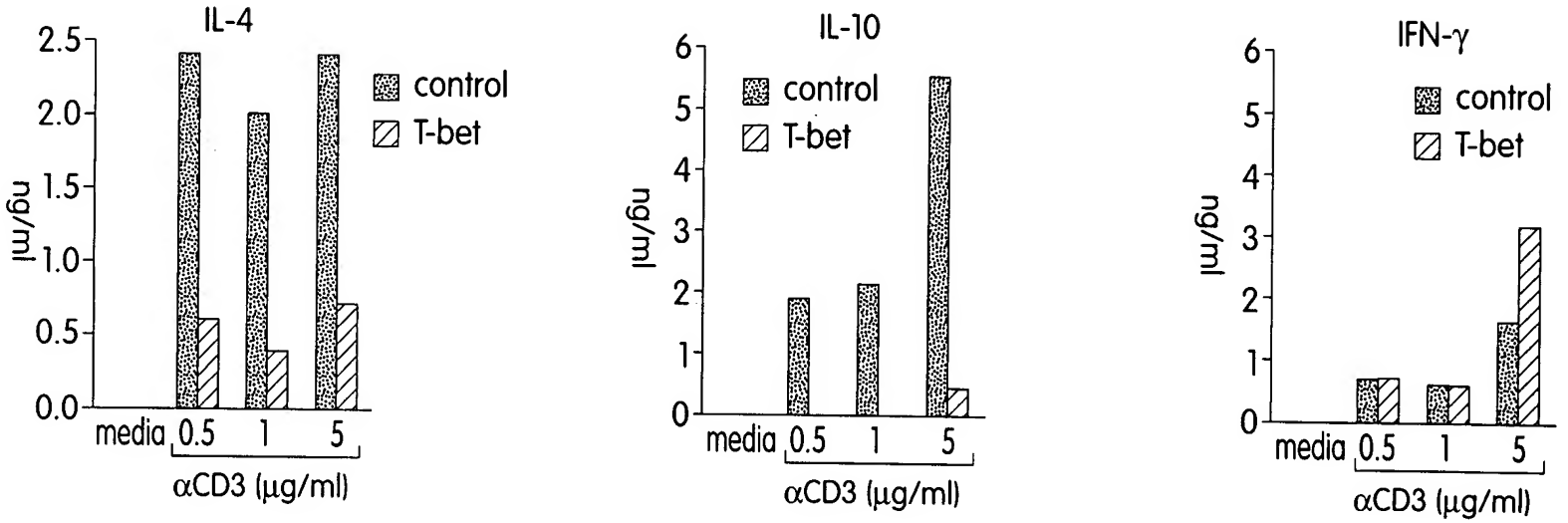
IL-4
2.5
2.0
1.5
1.0
0.5
0.0
ng/ml
control
T-bet
media
0.5
1
5
αCD3 (μg/ml)
IL-10
6
5
4
3
2
1
0
ng/ml
control
T-bet
media
0.5
1
5
αCD3 (μg/ml)
IFN-γ
6
5
4
3
2
1
0
ng/ml
control
T-bet
media
0.5
1
5
αCD3 (μg/ml)

Fig. 6

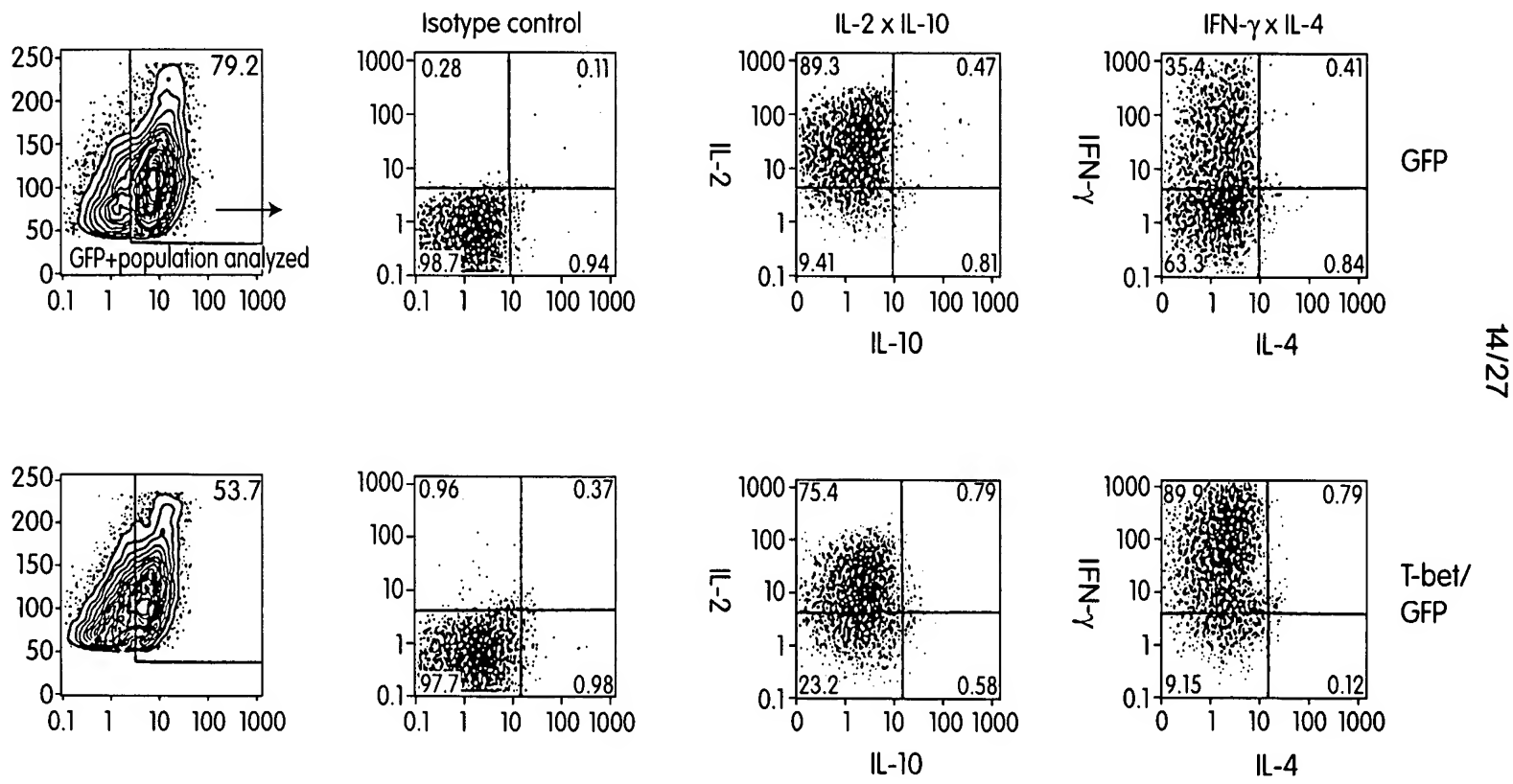
Isotype control
IL-2 x IL-10
IFN-γ x IL-4
79.2
GFP+population analyzed
0.28
0.11
98.7
0.94
89.3
0.47
9.41
0.81
35.4
0.41
63.3
0.84
GFP
IL-2
IL-10
IFN-γ
IL-4
53.7
0.96
0.37
97.7
0.98
75.4
0.79
23.2
0.58
89.9
0.79
9.15
0.12
T-bet/
GFP

Fig. 7

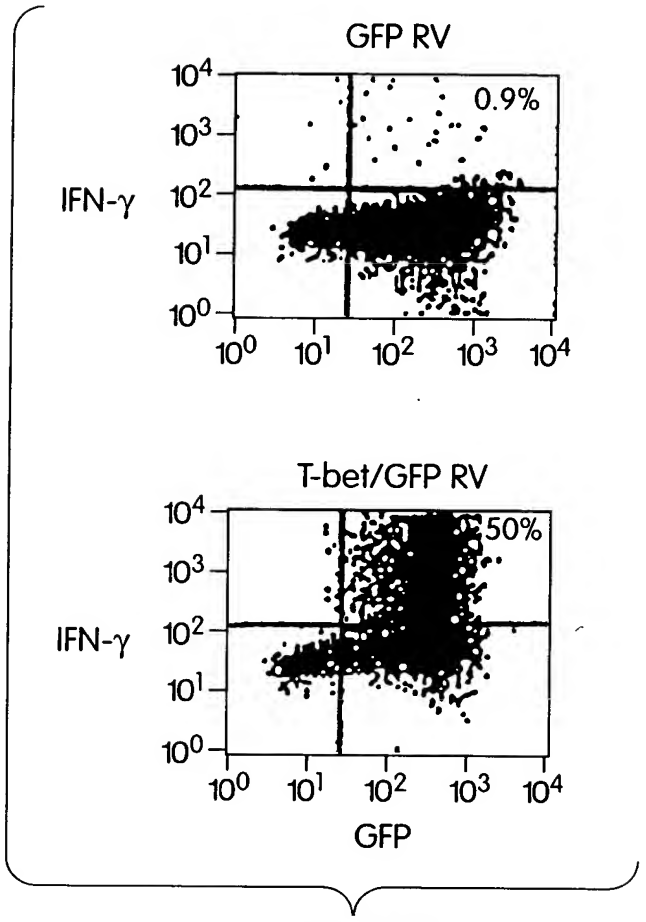
GFP RV
0.9%
IFN-γ
10^4
10^3
10^2
10^1
10^0
10^0 10^1 10^2 10^3 10^4
T-bet/GFP RV
50%
IFN-γ
GFP

Fig 8A

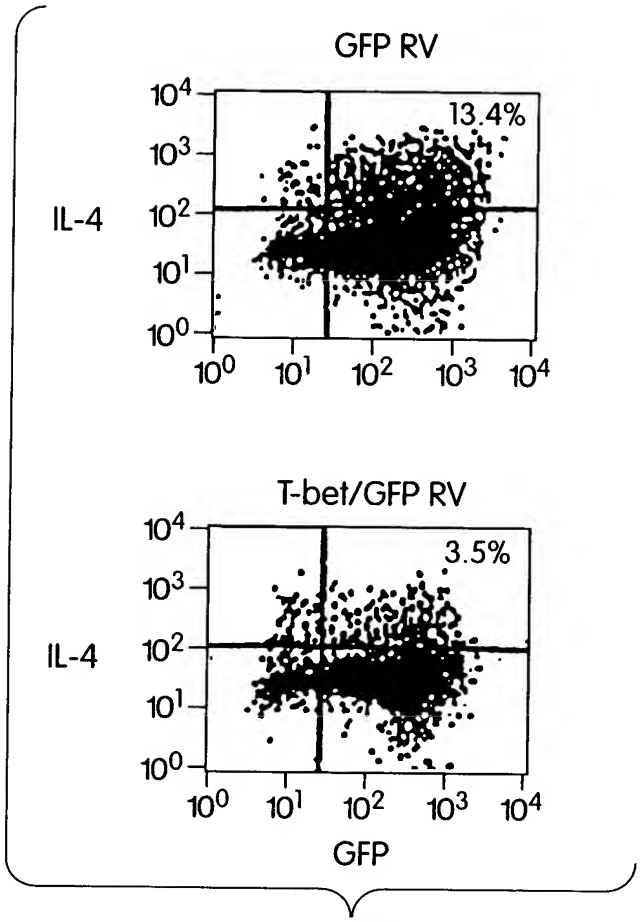
GFP RV
13.4%
IL-4
10^4
10^3
10^2
10^1
10^0
10^0 10^1 10^2 10^3 10^4
T-bet/GFP RV
3.5%
IL-4
10^4
10^3
10^2
10^1
10^0
10^0 10^1 10^2 10^3 10^4
GFP

Fig 8B

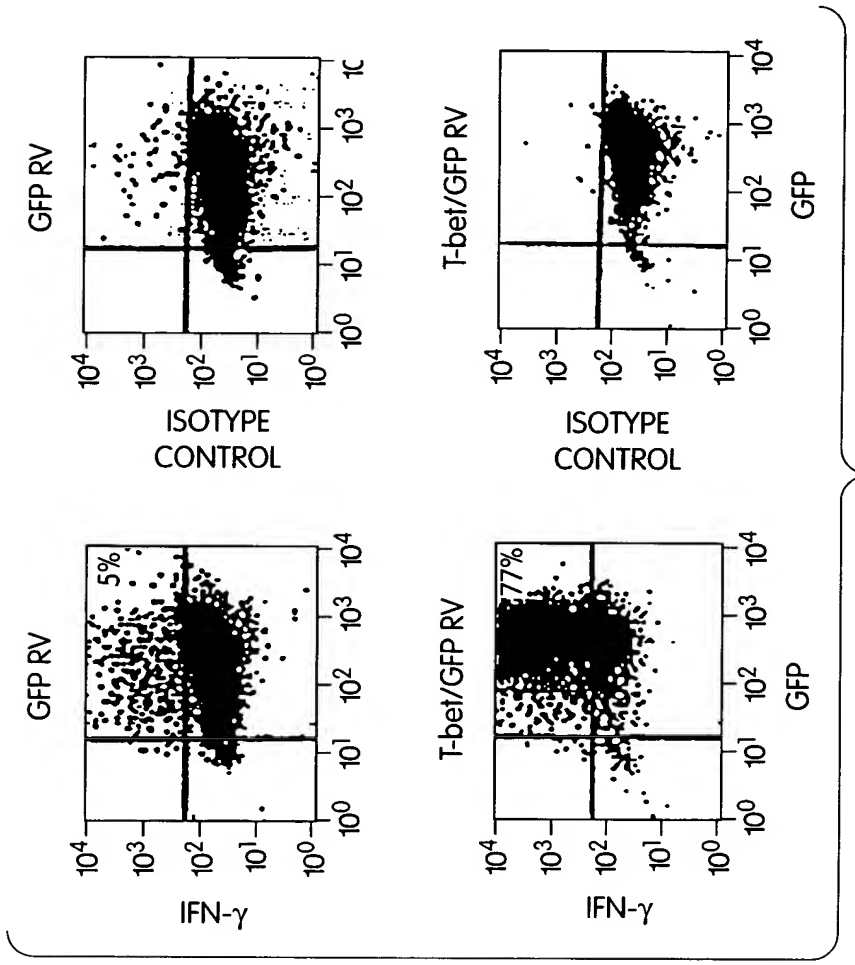
GFP RV
ISOTYPE CONTROL
T-bet/GFP RV
ISOTYPE CONTROL
GFP RV
5%
IFN-γ
T-bet/GFP RV
77%
IFN-γ
GFP
GFP

Fig 9A

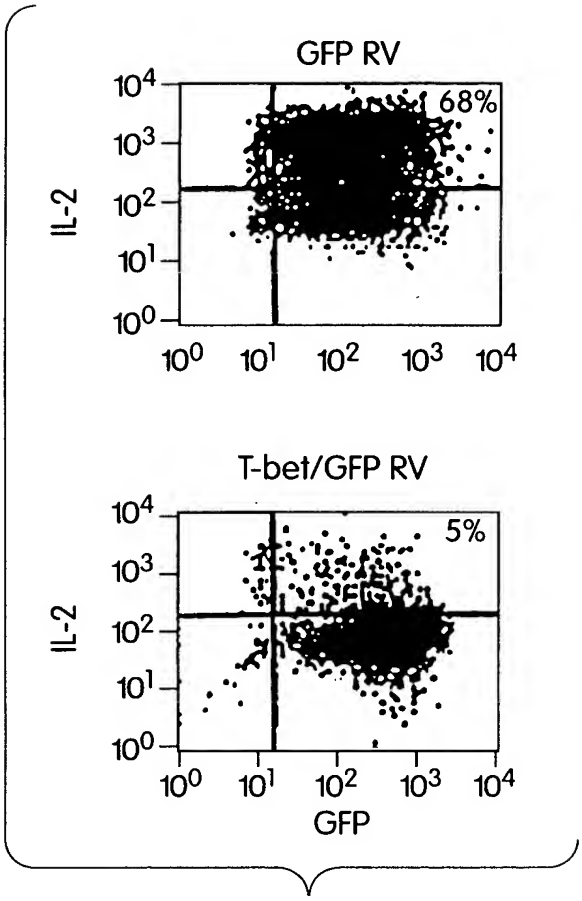
GFP RV
68%
IL-2
10^4
10^3
10^2
10^1
10^0
10^0 10^1 10^2 10^3 10^4
T-bet/GFP RV
5%
IL-2
10^4
10^3
10^2
10^1
10^0
10^0 10^1 10^2 10^3 10^4
GFP

Fig 9B

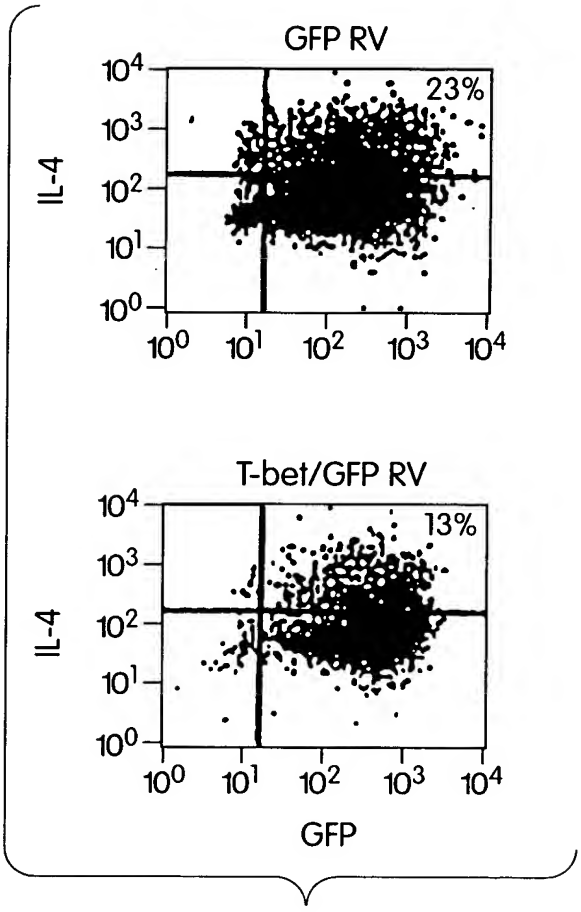
GFP RV
23%
IL-4
10^4
10^3
10^2
10^1
10^0
10^0 10^1 10^2 10^3 10^4
T-bet/GFP RV
13%
IL-4
10^4
10^3
10^2
10^1
10^0
10^0 10^1 10^2 10^3 10^4
GFP

Fig 9C

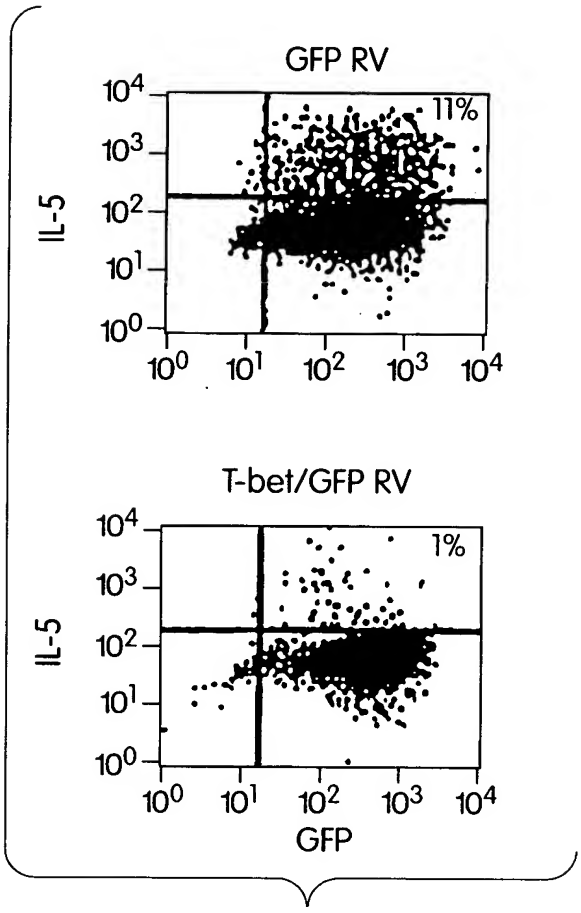
GFP RV
11%
IL-5
10^4
10^3
10^2
10^1
10^0
10^0 10^1 10^2 10^3 10^4
T-bet/GFP RV
1%
IL-5
10^4
10^3
10^2
10^1
10^0
10^0 10^1 10^2 10^3 10^4
GFP

Fig 9D

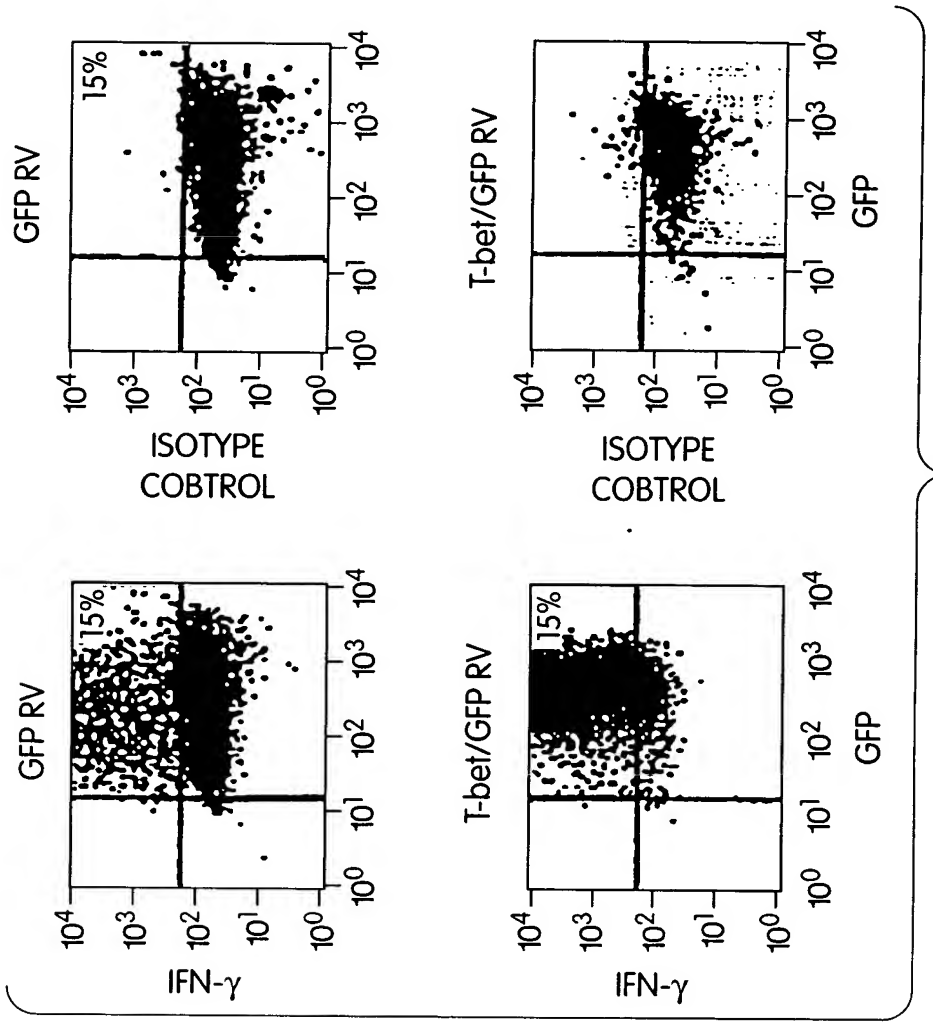
GFP RV
15%
ISOTYPE
COBTROL
T-bet/GFP RV
GFP
IFN-γ
10^0
10^1
10^2
10^3
10^4

Fig 10A

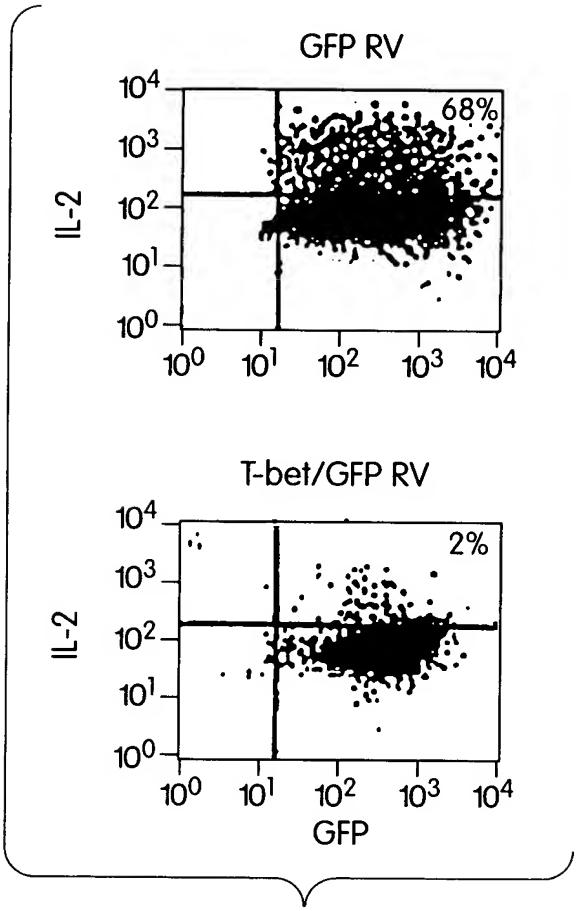
GFP RV
68%
IL-2
10^4
10^3
10^2
10^1
10^0
10^0 10^1 10^2 10^3 10^4
T-bet/GFP RV
2%
IL-2
10^4
10^3
10^2
10^1
10^0
10^0 10^1 10^2 10^3 10^4
GFP

Fig. 10B

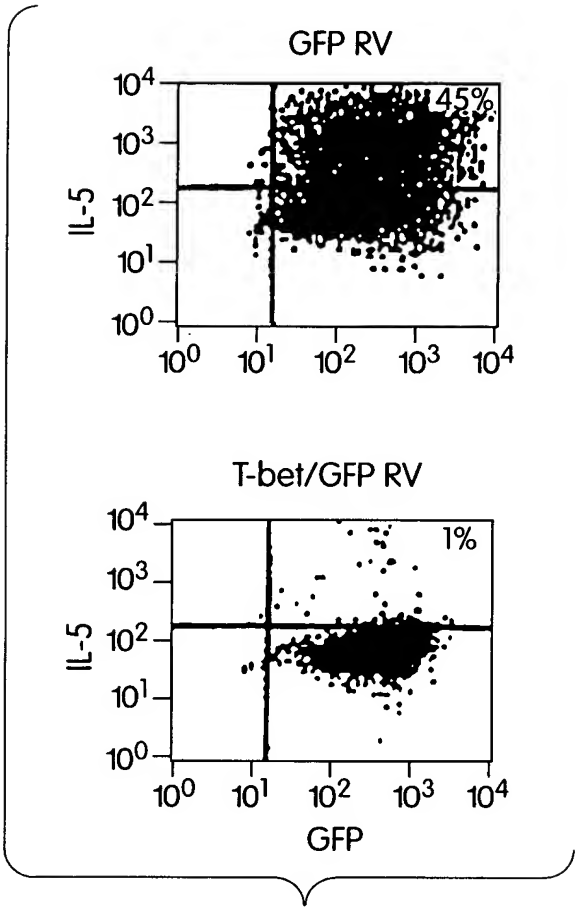
GFP RV
45%
IL-5
10^4
10^3
10^2
10^1
10^0
10^0 10^1 10^2 10^3 10^4
T-bet/GFP RV
1%
IL-5
10^4
10^3
10^2
10^1
10^0
10^0 10^1 10^2 10^3 10^4
GFP

Fig. 10C

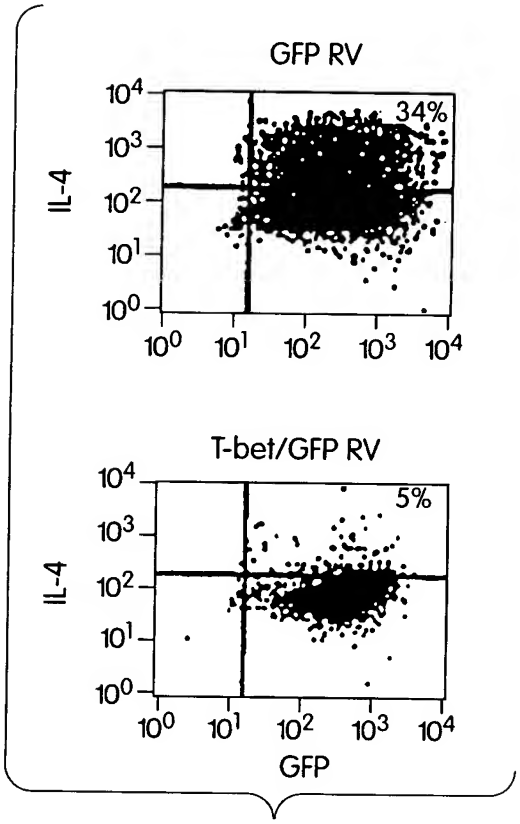
GFP RV
34%
IL-4
10^4
10^3
10^2
10^1
10^0
10^0 10^1 10^2 10^3 10^4
T-bet/GFP RV
5%
IL-4
10^4
10^3
10^2
10^1
10^0
10^0 10^1 10^2 10^3 10^4
GFP

Fig. 10D

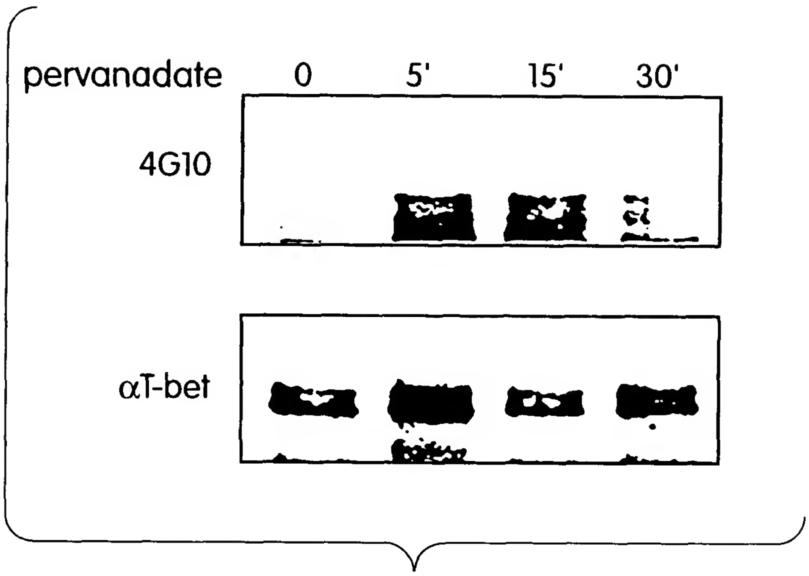

Fig 11

Continuation-in-part Application entitled:
T-BET COMPOSITIONS AND METHODS OF USE THEREOF

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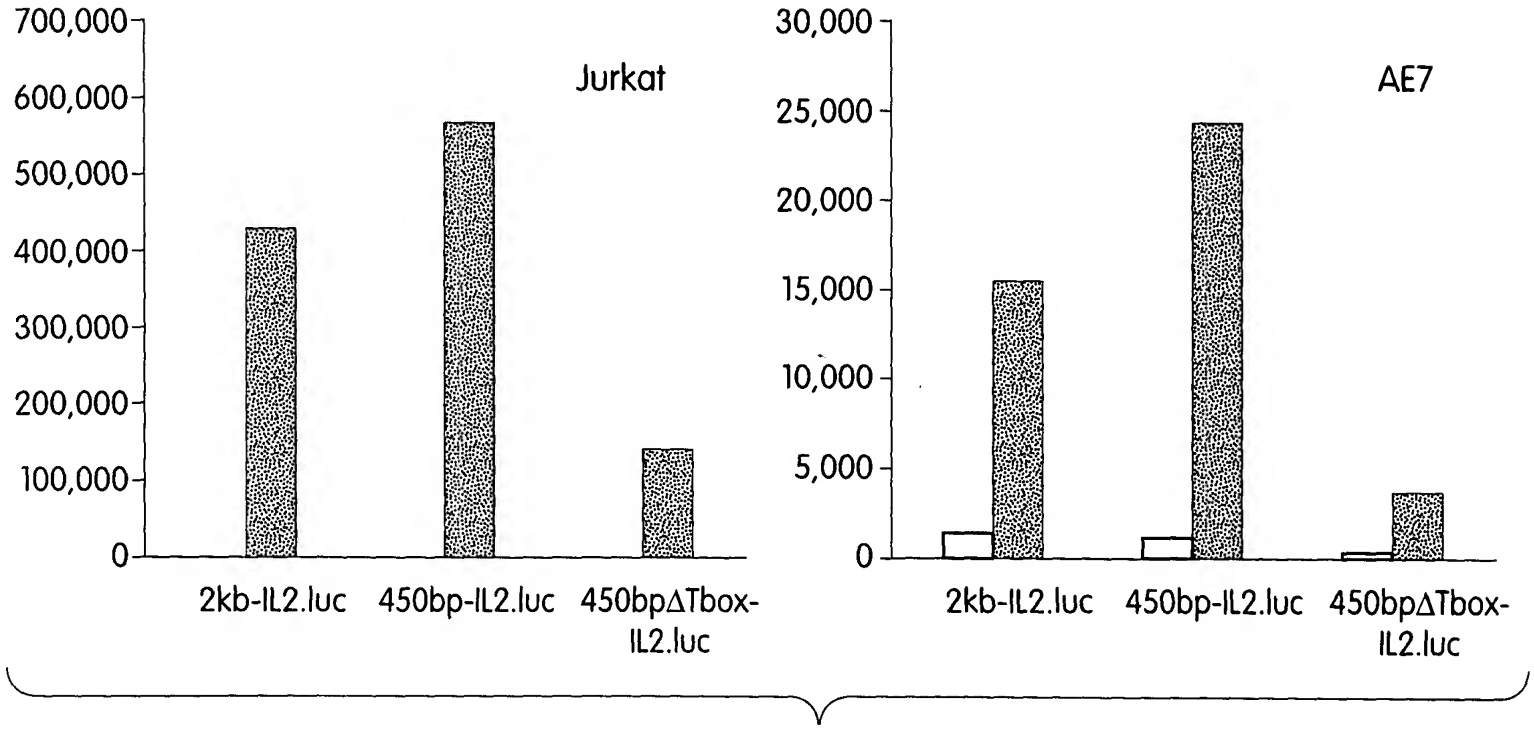
700,000
600,000
500,000
400,000
300,000
200,000
100,000
0
Jurkat
2kb-IL2.luc
450bp-IL2.luc
450bpΔTbox-IL2.luc
30,000
25,000
20,000
15,000
10,000
5,000
0
AE7
2kb-IL2.luc
450bp-IL2.luc
450bpΔTbox-IL2.luc

Fig. 13